

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
8 February 2001 (08.02.2001)

PCT

(10) International Publication Number
WO 01/08732 A1

(51) International Patent Classification⁷: A61M 11/00, (74) Agents: ELWE, Magnus et al.; Pharmacia & Upjohn AB, A61F 9/00, B05B 9/08, B65D 83/00 S-112 87 Stockholm (SE).

(21) International Application Number: PCT/SE00/01514

(81) Designated States (national): AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.

(22) International Filing Date: 20 July 2000 (20.07.2000)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
9902832-6 3 August 1999 (03.08.1999) SE
60/171,407 21 December 1999 (21.12.1999) US

(71) Applicant (for all designated States except US): PHARMACIA & UPJOHN AB [SE/SE]; S-112 87 Stockholm (SE).

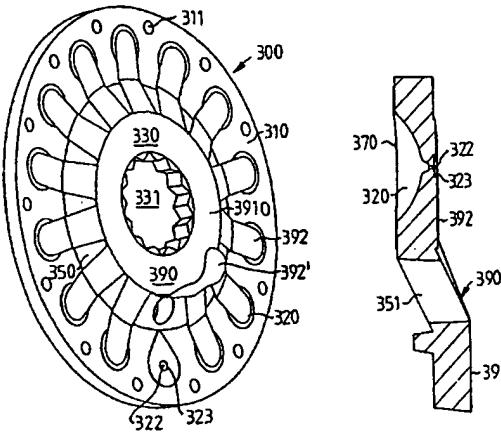
(84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published:

— With international search report.

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: LIQUID DELIVERY CONTAINER



WO 01/08732 A1

(57) Abstract: Pressurisable container for storing and ejecting liquid, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid. The front wall is substantially rigid in relation to the rear wall, the rear wall before pressurizing the container is substantially flat or substantially single-curved and the rear wall is deformable under stretching to substantially fill out the container cavity. The invention also relates to methods for container manufacture and devices and methods for ejecting liquid from the containers.

LIQUID DELIVERY CONTAINER

Technical field

The present invention relates to Pressurisable containers for storing and ejecting liquid, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid. The invention also relates to methods for container manufacture and devices and methods for ejecting liquid from the containers.

Background

Liquid containers designed not only to hold and store the liquid but also to deliver or expel the liquid tend to be growingly complex when the control demands on the delivery pattern increases. Whereas the complexity and expense, e.g. in pump systems, can be accepted under certain circumstances, such as in re-usable or multi-dosing devices, these conditions are not always present. It is for example often desirable to provide a single sealed container for each liquid dose to be delivered, e.g. to exactly control the dose and maintain sterility until the use moment in medical delivery applications, and under these unit dose circumstances the price restrictions becomes decisive. The design constraints may become still more severe with added requirements on delivery quality, e.g. in respect of delivery pressure, liquid speed, precise targeting, jet coherence, rapid stream rise and fall, fast delivery, small losses, precise dosing etc. High chamber pressures, e.g. to give high jet speeds or atomizing degrees, may counteract the cost aspect by requiring thick walls of special design or elaborate supports for the chamber or counteract the targeting by rupture, instability or dislocation of the opening. Jet coherence may require a precise opening channel inconsistent with minimum material and manufacturing conditions. High dosing precision requires complete chamber emptying and small losses, in turn requiring rapid pressure build up and fall, placing high demands on cavity stability and controlled collapse of the pressurizing wall. Secondary factors also need consid-

eration. It is for example common to provide a temporary seal over the opening in order to fully seal the container before use and arrangements need to be present for rupture or removal of the seal in connection with delivery. Also, apart from the container properties as such, certain rigidity and additional structures may be needed for retaining the container in a delivery 5 device fixture or seat. In order to avoid handling of individual unit dose containers it is also desirable to provide units of multiple connected containers for sequential firing in a dispenser device, which may require additional structural rigidity and features for feeding and stabilizing the individual containers in a device firing position. Manufacturing demands include both efficient production of the container parts as well as rational filling and sealing of the containers under high purity and even sterile conditions. 10

Prior art suggestions have met the abovesaid requirements only to a limited extent. The US patent specification 4090642 describes a tape having multiple pockets for a flowable material and represents liquid dispensing with low delivery and control requirements since the flowable material is only to be squeezed out on the tape surface for contact application to the 15 skin. The US 5497763 specification relates to a system including a similar multiple dose tape for atomizing liquid to be inhaled. The delivery requirements are still low as atomization takes place with a separate vibrator and the demands on the tape reduce to liquid outflow through a porous membrane. The GB 2255918 specification similarly relates to droplet formation for inhalation purposes but liquid atomization here takes place by forcing the liquid through narrow 20 container openings. In spite of the higher demands the proposed containers are separate collapsible containers or blister type sheets with multiple containers, requiring heavy support in the dispensing device when the pressure is applied to a dome shaped rear wall, the collapse of which cannot be fully controlled. It should also be noted that for the purposes described no liquid targeting is needed and no liquid stream formation since in connection with inhalation a 25 mist of droplets is passively drawn into the lungs by the patient, even allowing redirection of the stream. The same applies for powder inhalers, as exemplified by WO 90/13328, GB 2242134, DE 19500726, WO 97/04827, US 4811731, US 5207217, US 5415162, EP 469814, EP 129985 and US 4627432, additionally different in that no atomization at all takes place and no discharge from nozzle type openings. Accordingly such demands are not considered 30 and no container design suitable for such purposes is proposed. The WO 96/00050 and EP 224352 specifications do relate to the generation of a stream of droplets, actively shot and

targeted towards an eye and able to traverse an air gap by own inertia. Typically, however, the dispenser arrangements required are elaborate and no suggestions are given for individual unit dose containers with integral delivery nozzles. The WO 96/06581 and 97/23177 specifications suggest such unit dose containers for similar eye treatment purposes. While meritorious in 5 many respects the container designs proposed are similar to those already known and discussed, i.e. either elaborate separate containers or flexible bands with multiple blister type pockets having bulb type walls to be collapsed.

Accordingly there remains a need for improved integrated unit dose sealed container systems suitable both for liquid storage and liquid delivery under high control and quality 10 demands.

Summary of invention

A main object of the present invention is to avoid the abovesaid disadvantages of hitherto used container systems. A more specific object is to offer a unit dose sealed container system suitable both for liquid storage and liquid delivery under high control and quality demands. A further object is to provide such a system meeting high demands in respect of liquid 15 speed, precise targeting, jet coherence, rapid stream rise and fall, fast delivery, small losses, precise dosing and/or complete container emptying. Another object is to offer a system allowing high chamber pressures with maintained container integrity and without rupture, instability or dislocation of the opening. Still another object is to offer a system allowing controlled 20 collapse of the pressurizing wall. Yet another object is to offer a system useful for varying degrees of liquid stream momentum, e.g. characterized as low, moderate and high, allowing all from smooth application to penetrating strength. Another object is to offer system containers of low cost in material, component manufacture and filling. Still another object is to offer containers of high rigidity and stability. Yet another object is to provide such containers 25 suitable for multiple container units useful for sequential delivery of doses. Another object is to provide containers facilitating dispenser device design and requiring limited support at delivery. Another object is to provide a system easy to handle and convenient to operate for the end user.

These objects are reached with the characteristics set forth in the appended patent 30 claims.

By use of a container sealing, collapsible, backing rear wall which before collapse have an overall planar or single-curved shape several precision related objects are reached. For each front wall overall shape the rear wall will have a minimum surface, giving volume and shape stability before collapse and a foreseeable and controlled collapse during pressurization of the container as no buckling, folding or crumpling of the wall is needed as in corresponding flattening or inversion wall deformations. A stretchable material in the wall will allow the wall to increase its surface during collapse, hereby avoiding rupture, permitting dynamic adaptation to any ram or hammer tool surface used for forced displacement of the wall and allowing the wall to conform to the opposite container cavity surface, all serving to give controlled pressure response, including initiation, continuation and fall, and precision in dosing and container emptying. Manufacturing advantages are reached among others in that the wall form facilitates its production, its sealing to the front wall in filling procedures and its joint attachment to several containers in multiple compartment units or packages. Handling, convenience and dispenser device design advantages are reached in that the initial wall shape may be used to give smooth rear wall surfaces and does not build on the boundaries of the material used for the vessel front wall design. Said front wall is relied upon for overall container static and dynamic rigidity. Contrary to blister type containers, the present vessel wall can be designed freely and needs not depend on the curvature of pressed structures for rigidity, can have a wall thickness at each location adapted to its functional object and needs not be locally weakened by stretch operations and can have an integral opening determined by design and not by manufacturing necessities. Wall stability is essential to volume control and the possibilities for consistent emptying. Wall and opening strength is essential to pressure resistance and the operative range of the system, reliable targeting and stable and coherent liquid jets. It also reduces the needs for extensive container support arrangements in the dispenser device and facilitates sequential feeding of containers into a shoot position. If the rigidity advantages are used to avoid bulb wall parts, dispenser container seat and feeding mechanism can be further simplified. Still the option is retained of providing multiple container units and is now not limited to tapes and other flexible structures but can with advantage be formed into self-bearing structures with improved handling and convenience characteristics, similar to that of individual containers. The vessel wall can for example be implemented as the resulting structure when forming the container cavity in an otherwise flat or single-curved plate, al-

lowing both single and multiple container units of high structural rigidity and simple outer form, to the benefit of manufacture, dispenser design and end user handling, allowing free cavity designs e.g. for suitable flow characteristics and device head design as well as allowing opening designs of any length and form with high dynamic stability. Similar advantages are 5 obtained if implemented with substantially constant vessel wall thickness or with increasing wall thickness when moving lateral out from the opening location of the wall. The vessel walls with opening in place can be manufactured as an integral, single material, structure e.g. by heat molding in common plastic tools, without need for stretch operations or separate nozzle forming steps, and are easily filled with liquid and sealed by the rear wall.

10 Further objects and advantages of the invention will be evident from the detailed description hereinbelow.

Definitions

As used herein the expression "single-curved" shall be understood as any form of a plane or surface obtainable from the same plane in flat form, without stretching or shrinking 15 any part thereof in any direction within the plane, i.e. with maintained total surface for both the whole plane and any part thereof. In contrast, a "double-curved" plane or surface can only be obtained from a flat plane if deformed by stretching or shrinking. A curvature shall be regarded "continuous" if the change in curvature is constant or has a smooth variation and shall be regarded as "discontinuous" if there is a sharp change in the plane direction. As physical 20 and non-limiting examples a cylinder surface or a corrugated surface are single curved as they can be formed from a non-elastic sheet, e.g. paper sheet, by pure bending whereas the surface of a sphere or saddle cannot be so formed without stretching and accordingly are double-curved. All these surfaces are continuous by having smooth curvature changes whereas for example a sheet with sharp folds are discontinuous at the fold lines.

25 As used herein "system" shall be understood to refer to the invention generally, when including its parts, such as the container and dispenser devices therefore, as well as methods for part operation and use.

In the absence of explicit statements to the contrary, as used herein expressions like "comprising", "including", "having", "with" and similar terminology shall not be understood 30 to be exclusively restricted to recited element but shall be understood to allow for the presence of further elements as well and shall be understood to cover any element in integral, subdi-

vided or aggregate forms. Similarly, expressions like "connected", "attached", "arranged", "applied", "between" and similar terminology shall not be understood to cover exclusively direct contact between the recited elements but shall be understood to allow for the presence of one or several intervening elements or structures. The same applies for similar expressions

5 when used for description of forces and actions.

Also as used herein, positional and directional statements for both the container and the delivery device, such as "axial", "front" and "rear" and "forward" and "rearward", shall be understood with reference to the liquid delivery direction, with respect to which a line centered in the container opening and drawn along the main or average delivery direction shall be

10 regarded as the system "axis" along which axis the liquid is delivered in the forward direction.

Detailed description

The container can be said roughly to include a front wall and a rear wall between which the container space, or enclosure, for the liquid is confined.

The container space is formed mainly between the front wall and the rear wall and includes a cavity part formed in, or surrounded by, the front wall. Although the space available for the liquid may be larger than the volume of the cavity proper, e.g. additional small volume in the opening and in a rearward protrusion or curvature of the rear wall, it is preferred that the cavity constitutes the major part of the space, especially when the rear wall in the preferred manner is substantially flat or single-curved. The cavity surface preferably has the shape of the space of a vessel, opening towards the rear wall and having its closed bottom toward the front wall. Among others in order to facilitate complete emptying and form adaptation between cavity and driving ram, it is preferred that the cavity has little, and preferably no, undercut parts when seen from the rear side. This may be accomplished with vessel surface parts being parallel with the cavity axis, e.g. cylindrical parts, but preferably the surface widens all the way from the axis and laterally outwards, to create a generally concave deepening when seen from the rear wall side. Although the surface may have sharp discontinuities, e.g. between an axial cylindrical surface and a more flat front part, it is preferred e.g. for rigidity reasons that the surface is substantially smooth and continuous. The cavity shape in cross-section planes perpendicular to the axis can have any shape, e.g. oval or polygonal, although preferably circular. Suitable overall forms are that of a dome or segment of a sphere. As known per se the space may be divided into two or more separate chambers, e.g. for sequen-

tial delivery or for mixing in connection with delivery, for example by use of interior walls running perpendicular or parallel with the axis. Such walls can be manually removable before actual delivery or may be rupturable in connection with delivery. Alternatively adjacent containers may contain different liquids and eject substantially simultaneously, either as two discrete jets or jets mixing in connection with ejection, e.g. by openings arranged offset and close to each other or directed towards each other.

The front wall is preferably more rigid than the rear wall, by comprising more rigid materials and/or material of thicker dimensions. For reasons outlined it is beneficial that the front wall provides the container and its opening with rigidity and stability and preferably the front wall alone is able to provide such rigidity, leaving the rear wall design free for other considerations. As long as these demands are met the front wall can have a variety of configurations. The front wall may have a roughly constant thickness when measured normal to the cavity surface, making the front wall shape about congruent with the cavity surface. It is preferred, however, that the wall thickness measured as stated has a variation and preferably so that the thickness increases when moving away from the axis. This gives the container as a whole a high targeting and pressure sustaining stability while allowing a short and low friction opening design. This in sharp contrast to blister type containers where stretch manufacture of the bulb results in weak lateral wall parts with low overall container stability. A preferred way of arranging for abovesaid variation in wall thickness is to make the front surface of the front wall substantially flat or single-curved, flat being most preferred, giving a lot of manufacturing and handling advantages as indicated. It is further preferred that also the rear surface of the front wall in the neighborhood of the cavity, and disregarding the form of the cavity proper, is single-curved and most preferably flat, giving similar manufacturing and handling advantages. Among others the flat or single-curved surfaces allow attachment thereto of originally flat sheets, films or foils without stretching thereof, e.g. a sealing film on the front surface and the backing rear wall on the rear surface of the front wall. Most preferably the front and rear surfaces are parallel or concentric respectively, giving e.g. overall plate or cylinder type structures, having good rigidity, handling and manufacturing properties.

The opening design can vary depending on the nature of the liquid stream to be produced, e.g. an atomized spray or a concentrated stream to remain coherent or to break up into a linear stream of discrete droplets. Also the stream speed may vary from high penetrating to

low impact surface delivery. Several or multiple openings can be provided, e.g. to produce a controlled shower, although for most applications a single opening is preferred. The opening geometry can be that of a simple tube, diverging, e.g. for assisting in a distributed spray, converging, e.g. for assisting in a coherent stream to be formed, or a combination, such as a venturi type of channel. It is generally preferred to make the duct part of the opening short in order to keep the flow friction low. The desirable front wall thickness at the opening area can still be controlled, e.g. for rigidity purposes, e.g. by cutouts around the opening, preferably on the front surface of the front wall, also assisting in attaching a sealing film over the opening in a manner not interfering with the opening proper. The positioning of the opening with respect to the cavity can be asymmetrical, for example laterally offset towards the cavity side e.g. for mixing of adjacent jets, although normally it is preferred that the opening is symmetrically arranged with respect to the cavity, e.g. concentric with any cavity symmetry.

Although the above considerations apply to individual containers for single and discrete use it is preferred to provide for multiple container units or packages. This can be done by joining several individual container into multiple structures, e.g. by flexible joints to allow structures that can be bent, folded or rolled. Preferably, however, the multiple container package is a substantially rigid and self-bearing structure, among others giving advantages in connection with the delivery device. A rigid structure can be obtained by joining the individual containers by rigid joints but a preferred way is to utilize the rigidity of the front wall as described by providing an enlarged front wall structure and provide several cavities in the structure, among others facilitating manufacture of multiple containers and allowing a smooth and non-complicated exterior. By utilizing the feature of making the front and/or rear surface of the front wall structure flat or single-curved the attachment of films over these surfaces is further simplified, especially if the container surfaces lies in the same plane since an undivided sheet material can then be attached to several, and preferably all, of the individual containers of the structure, e.g. a common foil attached as rear wall to the rear surface of structure or a common peel sheet over the container openings at the front surface.

The overall shape of the front wall structure for multiple containers can take a variety of forms. A single-curved form for example can take the form of a partial or complete cylinder, allowing many containers on a limited volume and giving a very rigid structure. The container openings can be arranged on the cylinder exterior, e.g. if the dispenser device ram is

arranged to hit from inner or concave side of the full or partial cylinder or if openings are provided diametrically opposite each rear wall for ram access therethrough from outside the cylinder. The container openings can be arranged on the cylinder interior, e.g. for simple exterior ram access to the rear wall and also in this case a diametrically opposite hole can be provided

5 for passage of the liquid jet. Essentially flat front wall structures give advantages in manufacture and delivery device design. The shape may for example be rectangular, square or round. The round "disc" shape has been found particularly beneficial, among others in connection with the delivery device where sequential feeding of the containers into a shoot position can be made by simple rotation in a "revolver" type manner, the absence of any particular start

10 position facilitates handling and counting arrangements and allows for self-centering properties.

The layout of the multiple containers on the available structure surface can be made in various ways. The containers can be arranged in a row or line, e.g. to avoid two dimension feeding. Alternatively several rows can be provided along several lines in two dimensions,

15 e.g. parallel or concentric, to allow for a compact container arrangement.

The above design considerations for the front wall mainly apply to the parts occupied by containers, but other structures may be included for secondary purposes, e.g. for gripping, holding, centering or feeding discrete or multiple containers, in manufacture, in the delivery device and by the user, for adhering or removing sealing film etc. and these parts can be de-

20 signed freely for their respective purposes.

The front wall is preferably made from suitable inert plastic materials such as polyethylene or polypropylene, possibly with reinforcing fillings of e.g. glass fibres, and can be manufactured in various ways, e.g. the cavities with opening can be made by machining in a raw material blank but the structures described can easily be formed in one step, e.g. by

25 pressing although preferably by injection molding since all structures, including the opening, can be obtained with a dividable molding tool. Although the front wall can be made of a laminate or other composite material a single material is generally sufficient.

As said the rear wall serves the purposes of sealing the rear part of the container before ejection of the liquid and allowing collapse of the cavity during ejection. The rear wall should

30 run at least partially perpendicularly relative the opening axis, i.e. forming a non-zero angle therebetween and preferably runs substantially perpendicularly thereto. The requirements on

the rear wall material for sealing purposes depend on the particular application and liquid involved but generally it is desirable that the wall material is highly impermeable. It is also desirable to avoid solvents and other volatile components, both in the wall material and when attaching the rear wall to the front wall and preferably these parts are joined by heat sealing

5 rather than by glue or adhesives. For the collapse purposes it is possible to have a rear wall material that breaks or ruptures when hit with the ram of the dispenser device, which, however, requires good sealing between ram and cavity. Hence it is preferred that the rear wall can be deformed without rupture, and most preferably be deformed to such an extent that it can fill out and conform to the inner surface of the cavity. The requirements for deformability can be

10 reduced if the rear wall film is folded, e.g. in a continuous corrugated manner or in a discontinuous folded manner, i.e. in both cases with maintained single-curved shape, e.g. along single or multiple parallel lines over each container, to allow for unfolding as part of the deformation. In order to obtain a most controlled and reproducible collapse, however, it is preferred that no additional folds are used but that the rear wall when attached has substantially the

15 same shape as the rear surface of the front wall. In any case the deformation during ejection will result in that the rear wall flat or single-curved shape is at least to some extent changed into a double-curved form, requiring some stretching of the material. The stretching and deformation can be elastic, as in a rubber sheet, e.g. if repeated use is intended, but is preferably at least partially and preferably substantially plastic in an irreversible manner, e.g. in order to

20 avoid reuse of disposable containers or to prevent aspiration of air or liquid remains into the container after use. The rear wall preferably has the form of a sheet, film or foil of even thickness, which preferably is small compared to the front wall. The sheet may consist of a single material but preferably a composite material is used to meet all the requirements indicated.

Most preferably the sheet is a laminate. Preferably the laminate comprises at least one impermeable and diffusion retarding layer, preferably a metal layer such as an aluminum layer, and

25 at least one stretchable layer allowing deformation, preferably a plastic layer, e.g. of polyethylene, optionally also a heat sealable layer if the plastic layer does not have this property. In such a laminate it is acceptable that the metal layer breaks during wall collapse as long as remaining layers provide sufficient resilience for the deformation described.

30 As indicated it is preferred to temporary seal the container opening or openings before actual ejection of the liquid in order to maintain a fully sealed container. The seal should be

broken or removed immediately before use. Although a manually or pressure breakable rupturable seal can be used it is often preferred to use a removable seal in order to avoid any particle release from the seal, to have a fully foreseeable dynamic behavior and allow use of more reliable thick or strong layers. Generally a seal can be formed integral with the front wall, e.g.

- 5 by molding so as to leave a membrane of material somewhere in the opening duct. Preferably, however, a separate peel layer is provided for removal prior to ejection, and preferably attached to the front surface of the front wall. Again it is preferred to avoid glue and adhesives and preferably some form of welding is used, as by ultrasonic or heat. In order to facilitate removal and interference with the opening area the sealing can be made to a limited area sur-10 rounding the opening. In multiple container structures it is preferred to make the layers individually removable for each container, e.g. by use of separate films, pre-cut films or separate tongues, e.g. in a star shape for a round disc. For the film material similar considerations apply as for the rear wall material, although the film need not be deformable by stretching and the demands on impermeability can be slightly reduced in view of the small opening area.

- 15 The parts described should be joined and liquid filled into the cavity to form a pre-filled sealed container, typically containing liquid for a single dose to be delivered. Although possible to fill the container cavity through the container opening, e.g. through a needle, the preferred way is to fill the cavity from the rear side of front wall before attachment of the rear wall. A useful procedure is then to adhere the sealing film over the opening or openings, possibly under welding action, filling liquid into the container cavity from the rear side and adhering the rear wall over the filled cavity, again possibly under welding action. Preferably these steps are made in the order mentioned. The invention contributes in several ways to the efficiency and simplicity of such a procedure, particularly in contrast to blister type manufacture and filling. In contrast to blister bulbs, flat sheets type materials can be used for the sealing film and the rear wall, allowing flat or single-curved adherence with equally simple flat or roll type tools and allowing adherence over several containers in multiple container structures and no stretch or other material deformation steps are necessary. Adapted welding action, e.g. heat, is applied individually to the sealing film and the rear wall respectively and directly to the adhering materials involved, whereas in blister manufacture sealing heat for front and rear 20 materials has to be applied through a peel layer, over-adhering the peel layer, prolonging the process, possibly affecting the liquid present, and increasing product tolerances.
- 25
- 30

A delivery or dispenser device for ejection of the liquid from the containers can generally be said to comprise a housing with a seat for the container or container structure, a ram movable in relation to the housing in a direction substantially axial to the container when in the seat, an actuator arrangement operative to drive the ram.

5 The housing shall be understood in broad sense and may take a variety of forms. The device housing represents the point of reference for the container position and the movements described, such as the ram used for collapse of the container rear wall and for forces applied by actuating means performing said movements, whereat the force is applied between the housing and the moving part. The minimum functional requirement is that the housing offers
10 a support or platform for the container and movable parts and the actuating means providing the movements and forces. As in common practice, however, it is preferred that the housing forms a container at least partly embracing the parts and preferably also to such an extent that only the features designed to be controlled or monitored by the operator are externally exposed, to give an overall convenient design to use.

15 The housing should contain a seat for a container or several containers, the minimum requirements on which is that at least the container to be emptied is kept fixed in relation to the ram, preferably so that the container axis and the movement axis for the ram are parallel and most preferably coaxial with respect to the ram part to hit the container rear wall. Preferably the seat should be able to accommodate containers with the characteristics described
20 herein, e.g. with the shapes and sizes exemplified. The seat preferably supports the container against forward forces from the ram and preferably also against some rearward and lateral forces. The seat preferably allows the entire rear wall surface over the cavity to be exposed to the ram and should also expose at least the opening or openings on the front side of the container not to obstruct the liquid stream, although the rigidity of the present containers do not
25 require any heavy support. Preferably the seat is also designed to allow easy exchange of discrete containers, or sequential movement of the individual containers of a multi-container structure, into the active position of the seat, e.g. by having a track in which the structure can be moved in one or two dimensions. In the preferred embodiment of containers placed in a circle, preferably on a disc shaped structure, it is suitable to rotate the disc around a central
30 disc axis to bring the containers into alignment with the active position, in a revolver type manner. For single and in particular multiple container arrangements it is desirable that guid-

ing arrangements are provided to secure good alignment with the ram axis in order to reach high delivery precision intended, e.g. structures provided in connection with each container on the package for cooperation with at least one corresponding locking structure on the housing, seat or preferably ram, arranged for interlocking at proper alignment. Locking therebetween

5 can with preference be associated with a signal to assist stop in correct position, e.g. tactile or audible signal in manual operation or a mechanical or electronically detectable signal in automatic operation. Additionally it is preferred to include a counting arrangement, again manual or automatic, mechanical or electronical, designed to keep track on the number of containers used or remaining and warning for or preventing reuse of already emptied contain-

10 ers.

The ram may include a ram head and piston arrangement for moving the ram head along the movement axis. Although it is possible to design the ram head non-congruent with the container cavity, e.g. for use with different cavity shapes or when relying on rear wall stretch properties for emptying, it is preferred to design it for complete fill-out of the cavity.

15 This can be done with a soft and adaptable ram head, e.g. for the purposes of being compatible with different cavity forms, to increase operation range or to obtain a certain emptying pattern, preferably to squeeze out the liquid from the peripheral cavity parts towards the central, axial, parts which can be done for example by making the soft ram head slightly shallower in shape than the shape of the cavity vessel form. For a single cavity form it is, how-

20 ever, preferred to make the ram head front surface substantially identical with the inner cavity surface or, expressed in another way, the rear surface of the front wall in the container space. The ram head can be surrounded by a support, e.g. a tube structure in which the ram head travels, which is preferably also abutted around the cavity to seal the space between ram head and cavity at least during the rear wall collapse movement, e.g. to allow high pressures or re-

25 duce leakage risks. The piston part of the ram is generally not critical to the dynamics of the ejection but rather for propulsion and will be described in connection with the actuator system.

The ram can be propelled by use of a variety of mechanisms and energy sources. The mechanism can be operated directly with manual energy, in which case, however, it is pre-

30 ferred to provide an leverage or gear exchange to amplify or transform force or speed, preferably towards lower speed and higher force. In order to have controlled and consistent results

it is generally preferred to have automatic function in the sense that after operator triggering the propulsion takes place automatically, and preferably irreversibly, by use of stored energy. The energy may be stored in any way, e.g. in a mechanical spring, a gas spring or gas generator, as electrical energy or a combination thereof. The energy may be transmitted to the ram

5 by suitable motor or transmission means, e.g. electric motor or solenoid type motor for electrical energy, a piston and cylinder arrangement for gas springs or gas generators and rotation axis or plunger for coiled and helical springs respectively. It is generally preferred to include a transmission between the motor means and the ram proper, among other to provide a force amplification, e.g. by use of a gear wheel or a cam surface type of transmission. It is preferred

10 that at least the ram head, and preferably parts of the ram piston, are prevented from rotation during forward movement, which can be secured by any known guiding structures, such as a non rotation symmetric part cooperation with a complementary part, the parts being positioned on ram and housing respectively. A preferred transmission component for propulsion of the ram is a screw and nut arrangement, one of which is positioned on the ram and the other

15 on the motor side of the transmission. The necessary speed, force and movement characteristics for the ram depends on a number of conditions, such as the nature of the container parts and opening, the particular application implementation, e.g. surface or penetrative delivery, the viscosity of the preparation, e.g. aqueous solutions or ointments, etc. and general statements cannot be given. However, the energy sources, motor means and transmissions exemplified can be adapted to each need. It has also been found beneficial to include a damper, e.g. a dash pot, a linear damper, a flow valve, a magnetic damper etc., to control speed with maintained stable force. In most applications it is desirable to have a rapid rise and fall of pressure, generally requiring a stable and non-retarded speed of the ram, which is facilitated e.g. by a damper or high inertia in ram and transmission.

25 It is also preferred to include in the device arrangements to facilitate breakage or removal of the temporary seal over the openings as described. Although it is possible to break a seal by the pressure itself generated when collapsing the rear wall it is preferred to use an active step to break the seal. This can be done by having a de-sealing tool arranged in connection with the housing, e.g. a penetrating tool for a rupturable sealing or a wedging or drawing

30 arrangement for removal of peelable sealing films. Such arrangements can be located at or close to the seat, e.g. to allow late action, or remote, e.g. if the seat area is crowded. The de-

sealing tool can be operated manually or automatically or compulsory, e.g. as a part of the movement of container into the seat site. It has been found beneficial, however, to position the de-sealing tool on the rear side of the container for movement from rear to front, which allows the de-sealing tool to attack the film in the best manner possible, i.e. on the film rear side to

5 lift it from the front wall front surface. It also allows the de-sealing tool and mechanism to be arranged more conveniently within the housing and to the rear of the container for less interference with the ejection area and ejection target. The tool can have a movement mechanism of its own but most preferably the tool is arranged on or in connection with the ram in such a manner that it moves together with the ram, utilizing the same movement mechanism and

10 facilitating a removal immediately before ejection and as an unavoidable part of the ejection procedure. Preferably the de-sealing tool passes though an opening in the front wall structure, and possibly also through the rear wall, at a location not occupied by the container cavity but covered by the sealing film. With preference the dimensions of such an opening and the de-sealing tool can be mutually adapted so as to act as a guiding arrangement, as described, for

15 final alignment of ram and cavity before activation. In operation the tool first lifts the film from the container opening and the ram head then hits the container rear wall. It is possible to perform these two steps in a single continuous movement for the ram, e.g. for simplest operation and latest possible de-sealing, or in a two-step operation, possibly requiring two triggering actions from the user, e.g. in order to enable the user to verify that the film has been properly removed. It might also be of interest to use different movements characteristics for the

20 two steps, e.g. a slow movement for the peel not to cause tearing or rupture and a fast action for ejection, which might require some shift arrangement, e.g. a gear shift, de-coupling of brake or damper etc. The general container design principles of the present invention strongly amplifies the above described advantages, among other by having a rigidity permitting front

25 wall use for guiding purposes and allowing areas outside the cavity part to be utilized without instability problems.

Depending on each application it may also be beneficial to equip the device with means assisting targeting and positioning. For example, when used for shooting liquid to the eye the device front with a cowling, eye piece or eye cup for abutment to the eye socket.

30 Penetrating applications may require small distance or direct contact between opening and target surface whereas larger surface treatment may require an end piece defining both an an-

gle and a distance. As known per se for dispenser devices in general the device may also include mechanical or electronic warning, alerting or timing means.

As indicated in the introduction the invention described herein may be used for a variety of purposes within and beyond the medical area and for any type of preparations, such as 5 chemicals, compositions or mixtures, in any container and delivered for any purpose. It may be used for liquids within a broad range of compositions, e.g. pure liquids, solutions, emulsions, dispersions, body fluids etc., and viscosities e.g. high viscosity ointments. For reasons outlined the system has certain special values in connection with medical delivery devices 10 where also the design constraints are more severe than in most other applications. For convenience the invention has been described in terms of this application.

A preferred use of the invention is in connection with ophthalmic treatment of the eye with medical. The common administration manner is by eye drops or ointments, however, having several disadvantages. Both methods generally deliver a substantially higher amount than can be absorbed by the eye, not only resulting in dosing uncertainty and loss of expensive treatment medical but also in potential side-effects when non-absorbed preparation is 15 drained away via the nasolacrimal duct, e.g. beta-blocking agents used in eye treatment has substantial systemic effects. Another problem is that the common administration methods tend to induce a blink reflex that may entirely destroy the treatment or at least introduce a high degree of uncertainty. Also the common methods do not provide an high degree of targeting 20 precision, e.g. ability to hit the iris part of the eye being the penetrable part of the eye for prostaglandin. The principles used for the present invention solves these problems, among others by the possibility to deliver small amounts of liquid, actively ejected and not determined by liquid surface tension, by the possibility of delivering the liquid with sufficient speed to beat the blink reflex and by the possibility to eject a concentrated and coherent 25 stream for precise targeting. Typical parameters for this application will be given below although the invention shall not be regarded as limited to any such exemplified parameter. A typical single dose volume for delivery to the eye can be less than 25 microliter, preferably less than 15 and most preferably less than 10 microliter. Generally the volume is at least 1, preferably at least 2 and most preferably at least 3 microliter. Since it is desirable that each 30 container contains a single unit dose, these figures also relate to the liquid volume charged and contained in the containers, possibly allowing for some overfilling to compensate for non-

ejectable amounts, such as liquid remaining as wetting film or in the container opening duct, e.g. 25% but preferably no more than 10% overfilling. In addition to the liquid the container may contain other material, notably gas such as air or a purging gas such as nitrogen or noble gases, e.g. to facilitate manufacture, assist in atomizing or act as pressure buffer, although in

5 many instances little or no gas need to be present. The container can for example have a maximum cavity diameter is about 1 to 20 mm, preferably between 2 and 10 mm, calculated as a circle equivalent surface if the cavity is not round. A suitable speed for the stream of drops or jet ejected should be a balance between on one hand enough linear momentum to traverse an air gap between opening and target, without gravity assistance, and to travel fast

10 enough not be obstructed by blinking and on the other hand not so fast as to cause inconvenient sensible impact on the eye. The ideal speed is to some extent dependent on the drop size used but as a general rule the drops should be able to traverse at least 1 cm, preferably at least 3 and most preferably at least 5 cm through air by own momentum, incorporating reasonable distances between opening and target. A suitable lower speed limit when leaving the opening

15 is 1, m/s, preferably at least 5 m/s and most preferably at least 10 m/s. Generally the speed is lower than 200 m/s and preferably lower than 100 m/s. A suitable drop size so defined should be sufficient not to be retarded too quickly and not to be easily redirected, e.g. to be inhaled, and preferably has a minimum diameter of 20 micron, preferably not less than 50 micron and most preferably at least 100 microns. Normally the size is less than 2000 micron and prefera-

20 bly less than 1500 micron. The stream may take the form of a shower or spray of atomized liquid droplets but preferably the stream is narrow and fairly coherent although even such a stream tend to break up into individual droplets after a certain time of distance. The above given values are intended to relate to spherical droplets and for multiple droplets to the weight average of particle diameters. A coherent stream tend to break up into droplets of a diameter

25 of roughly double the diameter of the stream. Accordingly suitable opening diameters for the containers are about half the above given drop diameters or roughly between 10 and 1000 microns, preferably between 20 and 800 microns. A suitable front wall thickness, from front of opening to rear wall, may range from 0,5 mm to 10 mm and preferably between 1 and 5 mm. The above considerations are fairly independent of liquid viscosity and tend to apply

30 both for solutions and ointments.

Figures 1A to 1G illustrates schematically in section various container designs and features useful in single or multiple container arrangements

Figure 2A, 2B and 2C show in rear flat view, diametrical section and enlarged detail a multiple container structure in disc form, having 14 containers distributed around the disc 5 periphery.

Figure 3A to 3D depicts a similar disc as in Figure 2 in perspective views and partial sections, showing also rear wall and peel film.

Figure 4A and 4B shows in partial section and perspective view respectively the internal parts of a dispeser device for disc type multiple containers.

10

Description of drawings

In Figure 1 all embodiments, except that depicted in figure 1F, the front wall is formed from a plate type structure, being either flat as in Figures 1A, 1B, 1D and 1E or curved with concentric surfaces as in Figures 1C and 1G. The Figures illustrates various useful features but, for clarity, not all necessary or preferred details are shown in all Figures.

15

In Figure 1A the container 110 comprises the front wall 111, having a front surface and a rear surface being substantially parallel. The front wall 111 has a cavity 112 in the form of a vessel with an front vessel surface 113 and an opposite rear opening 114. The cavity cross-section shown has the general form of a circle segment. An opening 115, defining a container axis 116, provides a fluid connection between the cavity interior and the exterior at 20 the front wall front side. The opening 115 is here shown as a substantially constant cross-section channel with a thin inner membrane 117, formed from material left at injection molding and acting as a temporary seal to be broken at cavity pressurization. A deformable rear wall 118 covers the rear opening 114 of the cavity vessel with a seal (not shown). The rear wall is formed from an originally flat sheet material, here shown with discontinuous curvature 25 in the form of a number of folds 119, running perpendicular to the Figure plane, the folds acting to reduce the necessary stretching of the rear wall material when forced into the container cavity.

Figure 1B shows a container 120 similarly designed as in Figure 1A except that the opening 125 is shown slightly convergent and is covered on the front wall front surface by a 30 temporary seal in the form of a rupturable or peelable sheet 127, sealed to the front surface of the front wall. The rear wall 128 is here shown as a film with a continuous curvature part 129,

providing an additional volume over that provided by the vessel when limited by the rear vessel opening 124, here illustrated with a dotted line flush with front wall rear surface, and also adding surface to the rear wall limiting stretching thereof. The curvature of rear wall part 129 can be formed from a flat sheet material if the curvature is constant normal to the Figure 5 plane, as for a cylinder mantle surface, but need a double-curved material if in the form of a bulb.

Figure 1C illustrates a container 130 formed in a front wall 131 basically in the form of a cylinder part with single-curved front and rear surfaces centered around a cylinder axis 133 and with an opening 135 on the cylinder exterior, convex, surface. Also illustrated is a 10 cavity 132, having a cylindrical part 134 terminating in a sphere segment part 137, both parts symmetrically centered around opening 135 axis 136. Rear wall 138 is formed from a single-curved material continuously curved to adapt to the cylindrical inner surface of the front wall 131, hereby being larger in surface than a corresponding flat sheet over the cavity vessel opening, illustrated by dotted line 139.

15 Figure 1D illustrates a container 140 with a front wall 141 having a cavity 142 the vessel surface of which has a cylindrical part 143 and a flat bottom part 144. Also illustrated is an opening 145 with a surrounding recess 146 on the front wall front surface, serving to avoid direct contact between the opening proper and a peel film 147 and its sealing surface 148, which is arranged in a ring outside the recess 146.

20 Figure 1E schematically illustrates a container 150 with a cavity in the form of a circle segment, as in Figures 1A and 1B, centered around circle midpoint 153. Construction radii lines 154 drawn from point 153 intersects the inner surface 155 of the cavity 152 vessel perpendicularly and the Figure illustrates that the wall thickness, when measured along these construction lines normal to the vessel surface, increases (solid parts of lines 154) when 25 moving laterally away from the cavity axis 156. In contrast, Figure 1F illustrates a container 160 embodiment wherein the cavity 162 is formed in a front wall 161 design giving a constant wall thickness when measures along construction lines 164 centered in point 163 in the same manner as in Figure 1E. This Figure also illustrates a longer opening 165 duct, having a convergent part 166 and a straight part 167 with roughly constant cross-section. For clarity, sectioning lines have been omitted in these Figures.

Figure 1G illustrates a container 170 formed in a front wall 171 basically in the form of a cylinder part, as in Figure 1C, with single-curved front and rear surfaces centered around a cylinder axis 173. However, in this embodiment the opening 175 is arranged on the cylinder interior, concave, surface. Rear wall 178 is formed from a single-curved material continuously curved to adapt to the cylindrical outer surface of the front wall 171, hereby again being larger in surface than a corresponding flat sheet over the cavity 172 vessel. Also a temporary seal film 177, covering the opening 175, is formed from a single-curved sheet material.

All container embodiments illustrated in Figure 1A to 1G can be used as single discrete containers or can be part of multiple container structures by being joined with similar or 10 different container types, indicated by undefined lateral continuations of the front walls illustrated.

Figure 2A, 2B and 2C show in rear flat view, diametrical section and enlarged detail a multiple container front wall structure in disc form, having 14 containers distributed around the disc periphery. The disc, generally designated 200, can be said to include a peripheral 15 front wall structure 210 with the containers, a slightly offset central auxiliary part 230 with actuating and holding arrangements and a transition part 250 with peeling and guiding arrangements. In the embodiment shown the plate like basic shape of the front wall structure 210 has a front side 211 and a rear side 212 and comprises 14 identical cavities 220 in the general form of circle segments, the rear rim 221 of which are slightly rounded. The cavity 20 openings 222 are slightly convergent and on the front side 211 of the front wall structure 210 a small recess 223 is provided for reasons earlier outlined. The auxiliary part 230 can be said to include a central axis hole 231 for bearing and actuation purposes with a keying surface 232 providing non-rotational symmetry to facilitate actuation. The hole 231 preferably is slightly conical and converging from rear to front in order to facilitate actuator insertion and 25 centering. Spacing ridges 233 serve to provide a gap between discs when stacked, e.g. in sterilization. The transition part 250 can be said to include bores 251, also slightly converging towards the front, serving to allow penetration of a de-sealing tool from the rear side towards the front side to lift off a temporary seal in the form of a peel film (not shown), the cooperation between the tool and the bore also acting as a guiding arrangement, centering the cavity 30 with respect to a ram head of a dispenser device. With the cavity arrangement shown a flat rear wall film (not shown) can be positioned over the cavity rear sides to seal the containers,

e.g. after filling. Each cavity can be sealed with individual film parts but it is also possible to use a single sheet for covering all cavities. Preferably such films are radially restricted to the front wall part 210 of the disc, leaving the transition part 250 and the auxiliary part 230 uncovered. A typical overall thickness of the front wall plate can be about 2 mm with a cavity 5 depth of about 1.5 mm.

Figure 3A to 3D shows a similar disc arrangement as in Figure 2. Figure 3A is a perspective view of the disc front side, Figure 3B a perspective view of the rear side, Figure 3C a section from disc periphery to center through a cavity before ejection and Figure 3D a similar section of the cavity after use. As in Figure 2 the disc has a peripheral front wall structure 310 with the containers, a slightly laterally offset central auxiliary part 330 with actuating and holding arrangements and a transition part 350 with peeling and guiding arrangements. All details common with Figure 2 will not be repeated. Also in this embodiment there are 14 identical cavities 320 in the general form of circle segments with cavity openings 322 with a front side recess 323. Holes 311 serves to facilitate handling of the disc, e.g. during manufacture and filling. The auxiliary part 330 here has a central axis hole 331 for bearing and actuation purposes with 14 symmetrically arranged teeth 332, allowing the disc to be attached to an correspondingly toothed actuating axis of a dispenser device without consideration to the positioning of any individual container or disc part. The transition part 350 has bores 351, serving to allow penetration of a de-sealing tool from the rear side towards the front side to lift off 10 a temporary seal in the form of a peel film to be described. A single flat rear wall film 370, cut into ring form, radially restricted to the front wall structure 310, covers all the cavity rear sides to seal the containers. As best seen in Figure 3A a peelable film 390 is attached to the disc front side so as to cover all cavity openings 322. The peel film is formed in star form 15 from a single flat sheet and comprises a unifying central ring part 391, concentric attached to the auxiliary disc part 330, from which ring part tongues 392 radiate, one for each cavity, so as to pass over the transition part 350 under covering of the bores 351 and further radially out to cover the cavity openings 322. Initially all cavity openings 322 are covered by the tongues 392. When the liquid from one of the container is to be dispensed a de-sealing tool (not 20 shown) is moved through the corresponding bore 351 from the disc rear side towards the front side, hereby lifting the corresponding tongue 392' to free and expose the corresponding cavity opening 322, as best seen in Figure 3A and 3D. The container content is then ejected by forcing 25

ing the rear wall 370 under stretching from the flat condition shown in Figure 3C to the cavity filling condition shown in Figure 3D.

Figure 4A and 4B illustrates a dispenser device for multiple container discs as described in relation to Figure 3. The device, generally designated 400, is shown without any 5 casing part of the housing. The device receives the disc 410 at its front side, to the right in the drawings. The device can be said to include a seat 420 for the disc incorporating a disc bearing axis 421, having teeth 422 corresponding to the teeth of disc central opening. The axis 421 has a bearing gear wheel 423 cooperating with a control gear wheel 424 connected to a thumb wheel 425 for sequential advancement of the individual containers into the ejection position to 10 be described. The thumb wheel can be actuated manually or a may be connected to a mechanism (not shown) for automatic incrementation in connection with the motor actuation cycle. A pawl and ratchet mechanism 426 secures that the disc occupies predetermined positions only and that it can be rotated in one direction only to avoid reuse of already emptied containers. These parts may also include counting means or other assistance (not shown) to urge for 15 disc replacement. In the Figures the lowermost disc part is in the active ejection position 427 where the seat may comprise additional support or guidance (not shown) for the disc, e.g. to against forward and possibly rearward movement for example to balance force from the ram. Preferably such support structures can be arranged on a door arrangement of the housing for inserting the disc. In this position the cavity is centered with respect to the ram, generally 20 designated 430. The ram includes a ram head 431 with a circle segment front form, adapted to the container cavity form. The ram head is part of a ram head support 432, also carrying a de-sealing and guiding tool in the form of a probe pin 433, arranged to penetrate bores, corresponding to the bores 351 in Figure 3, in the disc 410 for the described purpose of lifting a temporary seal film from the container opening and of centering and stabilizing the container 25 cavity with respect to the ram head 431. Probe pin 433 extends farther in the forward direction than ram head 431 in order to perform the above functions before the ram head hits the rear wall of the container. The ram further comprises an external screw-thread 434 for propelling the ram axially forwards and rearwards. The ram is keyed against rotational movements by guidance (not shown) of the ram head support 432 flat lower surface 435. The device further 30 comprises a motor and transmission system, generally designated 450 for actuation of the ram movements. A rotationally arranged but axially stationary drive nut 451 has an internal screw-

thread 452 for cooperation with ram external screw-thread 434 so as to move the ram axially at rotation of the drive nut. An angular viscous damper 453 has a stationary part 454 and a rotational part 455, connected to the drive nut 451 so as to rotate with the drive nut. The drive nut 451 has external gear teeth 456 by which the drive nut can be rotated. A rotationally arranged motor housing 460 has an external gear 461 of larger diameter than the drive nut 456. Inside motor housing 460 is arranged a motor spring 462, which can be cocked by rotation via a winding axis (not shown) entering through the open part 463, to the left in the drawings, of the motor housing 460. Rigidly attached to the motor housing 460 and concentric therewith is a catch wheel 464 with a rim groove 465 for cooperation with an arming pin 466 of a manually accessible safety switch 467. A fire system 470 incorporates a release system (not shown) with a fire button 471 having two activation positions, a first partial operation of the fire button allowing the motor and transmission system 450 to advance the ram to move the probe pin 433 into completed removal of the temporary seal film, but with no contact between container rear wall and ram head 431, and a second operation of the fire button 471 allowing the ram head 431 to complete its forward movement into the container cavity, as shown in the Figure. A typical propulsion speed for the ram during ejection can be for example about 2 mm movement performed during a time of about 50 msec, for a cavity depth of about 1.5 mm.

The invention is not limited to the embodiments described and exemplified but can be varied within the terms of the appended patent claims.

Claims

1. Pressurisable container for storing and ejecting liquid, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid, characterized in the improvement comprising,
 - 10 that the front wall is substantially rigid in relation to the rear wall,
 - that the rear wall before pressurizing the container is substantially flat or substantially single-curved and
 - that the rear wall is deformable under stretching to substantially fill out the container cavity.
- 15 2. The container of claim 1, characterized in that the cavity has the form of a generally concave deepening when seen from the rear wall side.
3. The container of claim 1, characterized in that the cavity has little, and preferably no, undercut parts when seen from the rear side.
4. The container of claim 1, characterized in that the front wall has a roughly constant thickness when measured normal to the cavity surface towards the front wall.
- 20 5. The container of claim 1, characterized in that the front wall has thickness, as measured normal to the cavity surface towards the front wall, increasing when moving away from the axis.
6. The container of claim 1, characterized in that the front surface of the front wall is substantially flat or substantially single-curved, at least in the area around the opening.
- 25 7. The container of claim 1, characterized in that the rear surface of the front wall is substantially flat or substantially single-curved, at least in the area around the cavity.
8. The container of claim 1, characterized in that the front and rear surfaces of the front wall in the neighborhood of the cavity, but disregarding cavity and opening as
- 30 such, are substantially parallel or concentric.

9. The container of claim 8, characterized in that the front wall has the overall shape of a plate or cylinder part.

10. The container of claim 1, characterized in that the opening duct has a cross-section which is one of roughly constant, roughly converging, roughly diverging or a combination thereof.

11. The container of claim 1, characterized in that the opening is designed to assist in atomizing the liquid.

12. The container of claim 1, characterized in that the opening is designed to assist in forming a coherent linear liquid stream.

10 13. The container of claim 1, characterized in that the front wall front side has a cut-out area around the opening.

14. The container of claim 1, characterized in that the container is connected to at least one other container to form a multiple container unit.

15 15. The container of claim 14, characterized in that the front wall surface of several containers lies in the same flat or single-curved plane.

16. The container of claim 15, characterized in that the front wall surfaces of several containers are covered by a single sheet material.

17. The container of claim 14, characterized in that the rear wall surface of several containers lies in the same flat or single-curved plane.

20 18. The container of claim 17, characterized in that the rear wall surfaces of several containers are covered by a single sheet material.

19. The container of claim 14, characterized in that the unit is a substantially rigid and self-bearing structure.

25 20. The container of claim 19, characterized in that the unit comprises an enlarged front wall structure in which several cavities with openings are provided to form the multiple containers.

21. The container of claim 20, characterized in that the front and rear surfaces of the front wall structure are substantially parallel in the neighborhood of the cavities, when disregarding the cavities and openings as such, to give a general plate form.

30 22. The container of claim 21, characterized in that the front wall structure has the overall shape of a disc.

23. The container of claim 22, characterized in that the several containers are positioned along at least one circle concentric with the disc periphery.

24. The container of claim 20, characterized in that the front and rear surfaces of the front wall structure are substantially single-curved and concentric in the neighborhood of the cavities, when disregarding the cavities and openings as such.

25. The container of claim 24, characterized in that the front wall structure has the overall shape of a full or partial cylinder.

26. The container of claim 25, characterized in that the several containers are positioned over two dimensions of the cylinder surface.

10 27. The container of claim 1, characterized in that the rear wall is folded in a continuous or discontinuous manner.

28. The container of claim 1, characterized in that the rear wall has substantially the same overall shape as the rear surface of the front wall.

15 29. The container of claim 1, characterized in that the rear wall is designed to be deformed elastically.

30. The container of claim 1, characterized in that the rear wall is designed to be deformed inelastically or permanently.

31. The container of claim 1, characterized in that the rear wall comprises a laminate.

20 32. The container of claim 1, characterized in that the rear wall comprises a metal layer.

33. The container of claim 1, characterized in that a temporary sealing is provided over the opening.

25 34. The container of claim 33, characterized in that the sealing is rupturable or removable.

35. The container of claim 33, characterized in that the sealing comprises a flat or single-curved sheet.

36. The container of claim 1, characterized in that the liquid volume is less than 25 microliter, preferably less than 15 and most preferably less than 10 microliter.

30 37. The container of claim 1, characterized in that the opening diameter is between 10 and 1000 micron, preferably between 20 and 800 micron.

38. The container of claim 1, characterized in that the font wall thickness is between 0,5 and 10 mm, preferably between 1 and 5 mm.

39. The container of claim 1, characterized in that the maximum cavity diameter is about 1 to 20 mm, preferably between 2 and 10 mm.

5 40. Pressurisable container for storing and ejecting liquid, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space 10 for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid, characterized in the improvement comprising,

15 that in the vicinity of the cavity the front wall has the overall shape, except for the cavity itself, of a flat or single-curved plate with substantially parallel or concentric front and rear surfaces,

that at least a part of the cavity is formed between the front and rear surfaces with the opening exposed on the front surface and the open part of the vessel exposed on the rear surface, and

that the rear wall being attached to the rear surface.

20 41. The container of claim 40, characterized in any characteristic of claims 1 to 39.

42. Pressurisable container for storing and ejecting liquid, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening 25 defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid, characterized in the improvement comprising,

30 that the front wall thickness, as measured along lines running through the cavity and normal to the vessel closed surface, increases when moving off-set from the container axis.

43. The container of claim 42, characterized in any characteristic of claims 1 to 39.

44. A method for manufacture of a container containing liquid, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid, characterized in the steps of,

forming a front wall with a cavity in the form of a vessel with an opening connecting the vessel with the front wall front surface,

introducing liquid into the vessel cavity, and

attaching and adhering a flat or single-curved rear wall film to the vessel cavity open part to enclose the liquid in the container.

45. The method of claim 44, characterized in the step of forming the front wall with cavity and opening by injection molding.

46. The method of claim 44, characterized in the step of adhering the rear wall film by welding.

20 47. The method of claim 46, characterized in the step of welding by heat.

48. The method of claim 44, characterized in the step of adhering a flat or single-curved sealing film over the opening.

49. The method of claim 44, characterized in the step of forming a front wall with more than one cavity.

25 50. The method of claim 49, characterized in the step of adhering the rear wall film over more than one cavity.

51. The method of claim 49, characterized in the step of adhering a flat or single-curved film over more than one cavity.

52. The method of claim 44, characterized in that the container has any of the 30 characteristics of claims 1 to 39.

53. A container containing liquid, characterized in that it is manufactured according to the method of any of claims 44 to 53.

54. A method for ejecting liquid from a container, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid, characterized in the steps of,

pressurizing the container by moving the rear wall at least partially in the axial direction and towards the opening with sufficient speed to eject liquid through the opening and hereunder stretching the rear wall, elastically or inelastically, to increase its surface.

55. The method of claim 54, characterized in that the stretching step comprises the step of stretching the rear wall from a flat or single-curved form into a double-curved form.

56. The method of claim 54, characterized in that the stretching step comprises the step of deforming the rear wall until substantially corresponding to the cavity form.

57. The method of claim 54, characterized in the step of substantially evacuating the liquid of the container.

58. The method of claim 54, characterized in that the liquid is ejected from the opening with a speed of at least 5, preferably at least 10 m/s.

59. The method of claim 54, characterized in that the liquid is ejected in the form of droplets of a diameter less than about 20 micron.

25 60. The method of claim 54, characterized in that the liquid is ejected in the form of a coherent jet.

61. The method of claim 54, characterized in that the liquid is allowed to pass through air a distance not less than 1 cm before hitting a target surface.

62. The method of claim 54, characterized in that the liquid is allowed to hit 30 an eye.

63. The method of claim 54, characterized in that the liquid is allowed to hit a soft surface for at least partial penetration thereof.

64. A device for ejecting liquid from a container, the container comprising a) a front wall having or surrounding a cavity corresponding to the form of an open vessel, b) an opening in the front wall adapted for ejection of the liquid from the container, said opening defining a container axis, c) optionally a sealing over the opening adapted for temporary use, and d) a rear wall closing and sealing the open part of the front wall vessel to confine a space for the liquid in the container, the rear wall running at least partially perpendicular to the container axis and being displaceable or deformable for movement towards the opening to pressurize the container liquid, characterized in the improvement comprising,

a housing with a seat for the container adapted to receive a container having a distance between rear wall and front wall front surface of at least 0,5 mm,

a ram arranged in a moving direction, in relation to the housing, substantially axial to the container when in the seat,

15 an actuator operative to drive the ram.

65. The device of claim 64, characterized in that the container when in the seat exposes substantially the whole part of the rear wall surface covering the cavity towards the ram.

66. The device of claim 64, characterized in that the seat is arranged to allow 20 exchange of containers in the seat.

67. The device of claim 66, characterized in that the seat is arranged to allow exchange by sequential feeding of containers in a multiple container unit into the seat.

68. The device of claim 67, characterized in that the seat comprises a track in which the containers can be fed.

25 69. The device of claim 67, characterized in that the seat allows sequential feeding by rotation of a multiple container unit having containers arranged in a circle pattern.

70. The device of claim 64, characterized in a guiding arrangement arranged to secure alignment between the ram and the container cavity.

71. The device of claim 70, characterized in that the guiding arrangement 30 comprises a releasable lock between the container and the housing or seat.

72. The device of claim 70, characterized in that the guiding arrangement comprises a releasable lock between the container and the ram.

73. The device of claim 72, characterized in that the locking arrangement comprises a structure locking the container when moved in the moving direction of the ram.

5 74. The device of claim 64, characterized in that the ram comprises a ram head and a ram piston.

75. The device of claim 74, characterized in that the front part of the ram head substantially conforms with the container cavity.

10 76. The device of claim 74, characterized in that at least the ram head front part is made of a soft material adaptable to the container cavity.

77. The device of claim 74, characterized in that the actuator is arranged to displace the ram piston.

78. The device of claim 64, characterized in that the actuator comprises an electrical arrangement for driving the ram.

15 79. The device of claim 64, characterized in that the actuator comprises a mechanical arrangement for driving the ram.

80. The device of claim 79, characterized in that the mechanical arrangement comprises at least one spring for energy storage.

20 81. The device of claim 64, characterized in that the actuator comprises a transmission including at least one driving force transforming arrangement.

82. The device of claim 81, characterized in that the transmission includes a screw and nut arrangement.

83. The device of claim 64, characterized in a damper arranged to affect the ram movement.

25 84. The device of claim 64, characterized in a de-sealing tool arranged for breakage or removal of a sealing over the container opening.

85. The device of claim 84, characterized in that the tool is arranged to the rear of the container when in the seat and arranged for forward movement during de-sealing.

30 86. The device of claim 85, characterized in that the tool is arranged to pass through or past the front wall during its forward movement to attack the sealing.

87. The device of claim 86, characterized in that the tool and container are arranged to cooperate as a guiding arrangement arranged to secure alignment between the ram and the container cavity.

88. The device of claim 85, characterized in that the tool is connected to the 5 ram for common movement therewith.

89. The device of claim 88, characterized in that the tool is connected to the ram so as to hit the sealing before the ram hits the container.

90. The device of claim 64, characterized in that the container is any of the containers of claims 1 to 39.

10 91. The device of claim 64, characterized in arranged to enable ejection of liquid in accordance with any of claims 54 to 63.

92. A kit or combination characterized in that it comprises a) a container according to any of claims 1 to 39 and b) a device having a ram arranged to displace or deform the container rear wall to pressurize the container liquid.

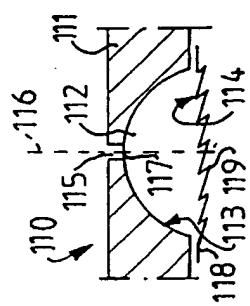


FIG. 1A

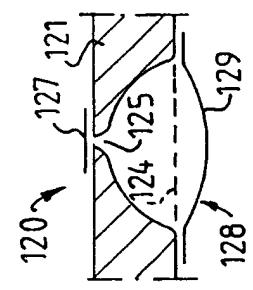
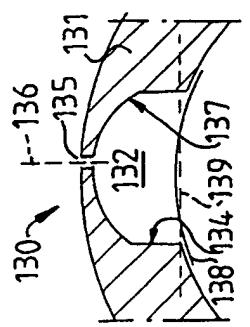


FIG. 1B



*133 FIG. 1C

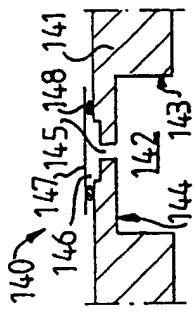


FIG. 1D

114

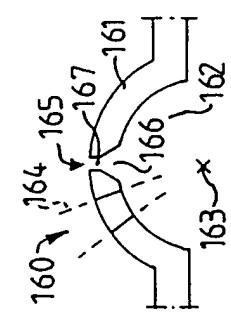


FIG. 1E

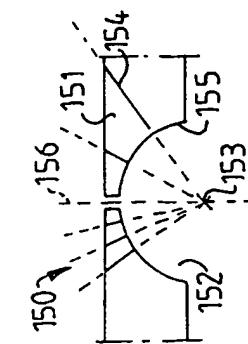
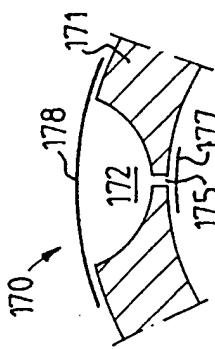
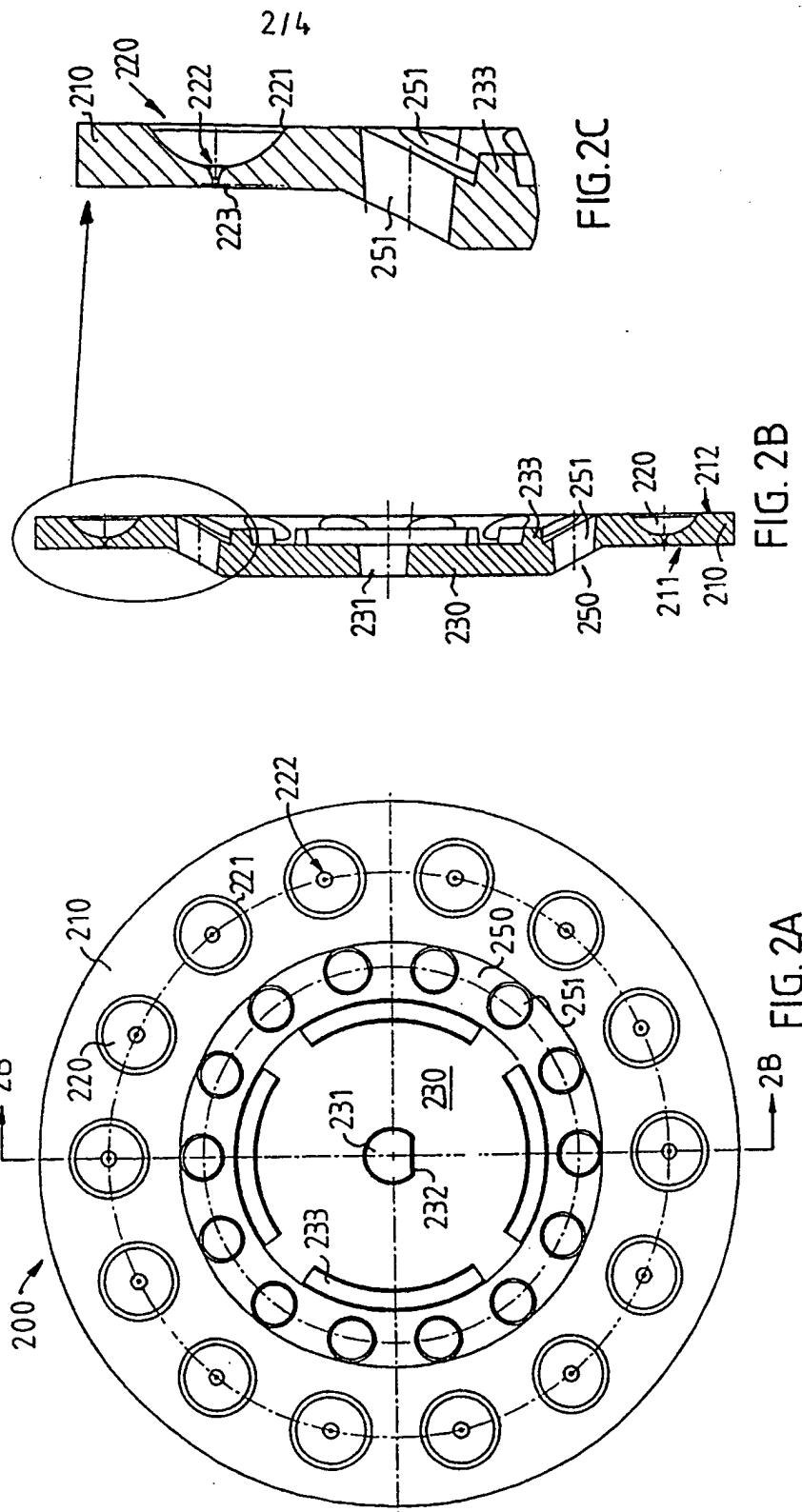


FIG. 1F



*173 FIG. 1G



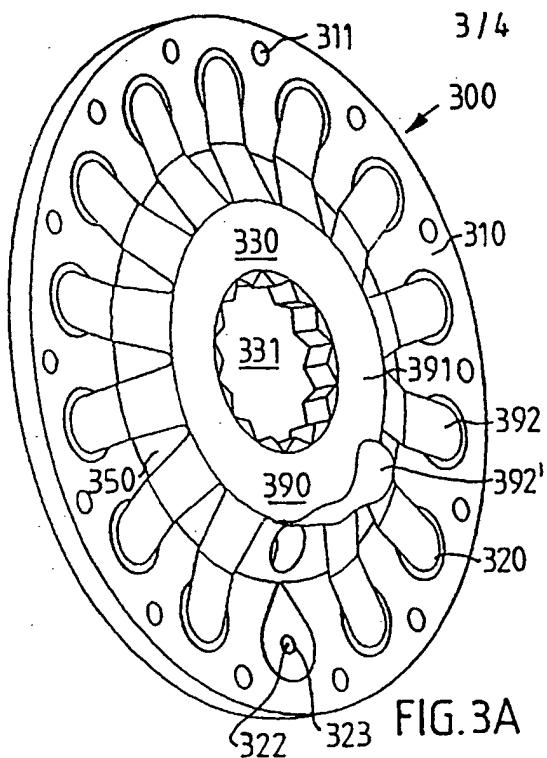


FIG. 3A

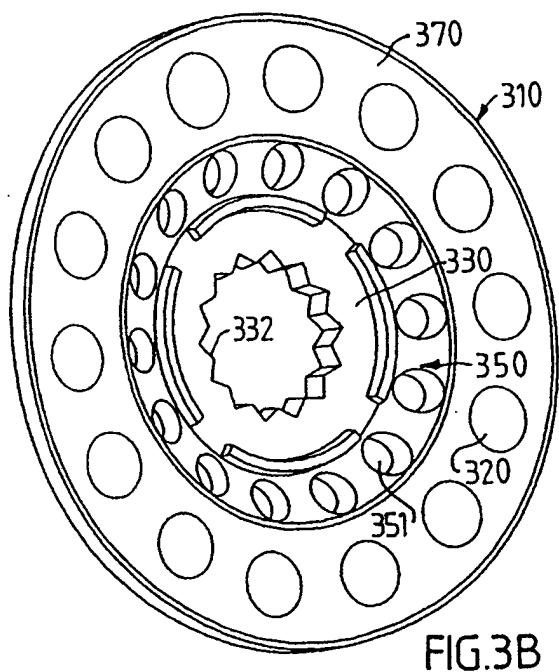
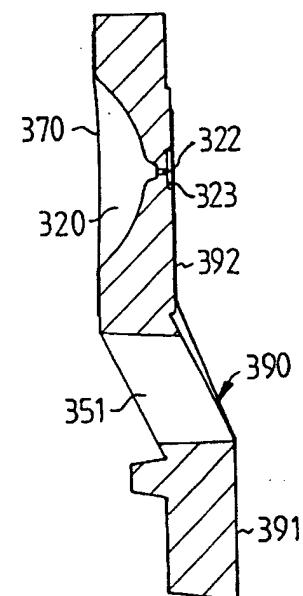


FIG.3B

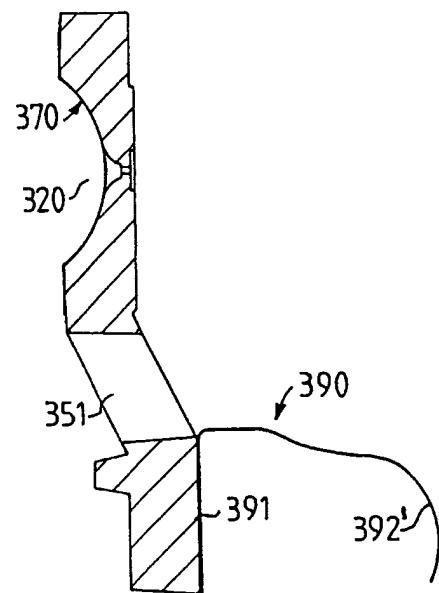
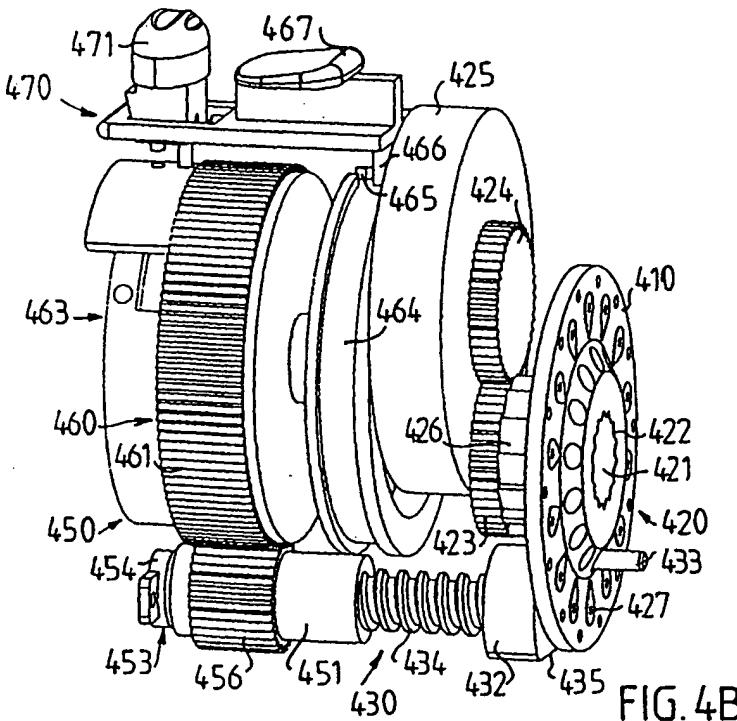
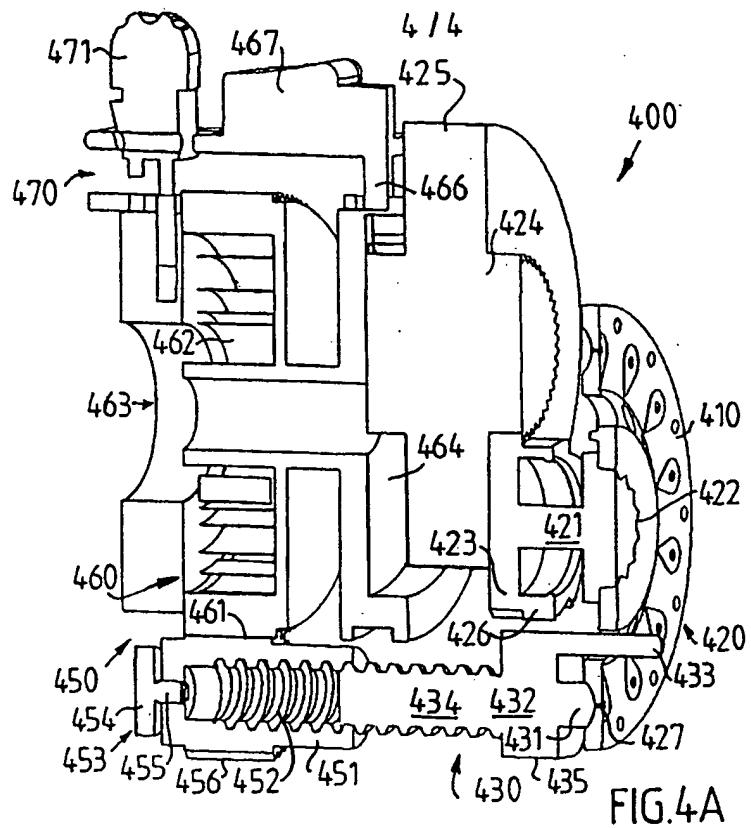


FIG. 3D



1
INTERNATIONAL SEARCH REPORTInternational application No.
PCT/SE 00/01514

A. CLASSIFICATION OF SUBJECT MATTER

IPC7: A61M 11/00, A61F 9/00, B05B 9/08, B65D 83/00
According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC7: A61F, B05B, B65D, A61M

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

SE,DK,FI,NO classes as above

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP 0832568 A2 (UNICE INTERNATIONAL LIMITED), 1 April 1998 (01.04.98), column 6, line 11 - line 34; column 7, line 11 - column 8, line 48, figures 1-3, claim 12, detail 18	1-4,6-8,10, 27-28,30, 33-35,38, 44-48,52-57
Y	--	5,9,29,41,43
X	EP 0139855 A1 (PLANTORGAN WERK HEINRICH G.E. CRISTENSEN KG), 8 May 1985 (08.05.85), figure 11, claims 6,9	1-4,7,28-29, 33-35,44, 52-57
Y	-- US 4433797 A (GALIA), 28 February 1984 (28.02.84), column 4, line 23 - line 27, figures 1-2, details 16,54	5,9,29,41,43
	--	

 Further documents are listed in the continuation of Box C. See patent family annex.

• Special categories of cited documents:	
"A" document defining the general state of the art which is not considered to be of particular relevance	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"E" earlier application or patent but published on or after the international filing date	"X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"O" document referring to an oral disclosure, use, exhibition or other means	"&" document member of the same patent family
"P" document published prior to the international filing date but later than the priority date claimed	

Date of the actual completion of the international search	Date of mailing of the international search report
9 November 2000	20 -11- 2000
Name and mailing address of the ISA/ Swedish Patent Office Box 5055, S-102 42 STOCKHOLM Facsimile No. +46 8 666 02 86	Authorized officer Marianne Dickman / JAn Telephone No. +46 8 782 25 00

INTERNATIONAL SEARCH REPORT

International application No.
PCT/SE 00/01514

C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 5709202 A (LLOYD ET AL), 20 January 1998 (20.01.98), column 20, line 20 - line 42; column 26, line 33 - line 39; column 27, line 28 - line 39, column 28, line 52 - line 59; figures 2-3,5-6, 9-10 --	40,42,64, 66-73,78-80, 84
Y	US 5516008 A (RABENAU ET AL), 14 May 1996 (14.05.96), column 3, line 56 - line 61; column 5, line 45 - line 55 --	11-18,31-32, 36-37,39, 49-51,58-63
X	WO 9606581 A1 (R.P. SCHERER CORPORATION), 7 March 1996 (07.03.96), page 18, line 4 - line 14; page 20, line 12 - page 21, line 5, figure 18 page 3, line 35 - page 4, line 2; page 15, line 10 - line 36, claim 1, detail 108 --	64-68,70-71, 74-77,79-80, 83
Y	WO 9723177 A1 (R.P. SCHERER LIMITED), 3 July 1997 (03.07.97), page 3, line 20 - line 21 --	11-18,31-32, 36-37,39, 49-51,58-63
A	GB 2255918 A (TECHNOLOGY LIMITED), 25 November 1992 (25.11.92), page 6, line 3 - page 7, line 31, figure 1, abstract --	1-18,27-48, 52-68,84, 90-92
A	US 4522622 A (PERRY ET AL), 11 June 1985 (11.06.85), figures 1-3 --	1,5,8-9, 14-23,40-43
A	WO 9600050 A1 (R.P. SCHERER CORPORATION), 4 January 1996 (04.01.96), figure 6, abstract --	1-3,27-43, 54-63

INTERNATIONAL SEARCH REPORT

International application No.
PCT/SE 00/01514

A US 1751341 A (M LEISSE), 18 March 1930 (18.03.30), 1,27,30
figures 1-3
--

A US 2649995 A (N. MUSKIN), 25 August 1953 27,30
(25.08.53), column 3, line 11 - line 19
--

A US 4236516 A (NILSON), 2 December 1980 (02.12.80), 1,54,64
figures 1-2
--

A US 5429280 A (BAUER ET AL), 4 July 1995 (04.07.95), 5,9,29,41,43
figure 8
--

A US 4072249 A (B. EKENSTAM ET AL), 7 February 1978 1,14,54
(07.02.78), column 4, line 32 - line 44; column 5,
line 50 - line 52; column 7, line 11 - line 14
--

A GB 2246555 A (CHRISTOPHER, G.), 5 February 1992 14
(05.02.92), abstract
--

A US 5622166 A (R.F. EISELE ET AL), 22 April 1997 14-23
(22.04.97), figures 2-4
--

A US 5349947 A (M.T. NEWHOUSE ET AL), 27 Sept 1994 14-20,24-25,
(27.09.94), column 9, line 30 - column 10, line 6, 49-51,64-77,
figures 5-12 79,81
--

A US 5415162 A (R.A. CASPER ET AL), 16 May 1995 24-26
(16.05.95), figure 4
--

INTERNATIONAL SEARCH REPORT

International application No.
PCT/SE 00/01514

A US 5221050 A (A. JEFFRIES ET AL), 22 June 1993 64-66,74-76
(22.06.93), column 1, line 29 - line 32; column 5,
line 4 - line 40

INTERNATIONAL SEARCH REPORTInternational application No.
PCT/SE00/01514**Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)**

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:

2. Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:

3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

Invention 1
Claims 1-13, 27-48, 52-63, 90-92. Relates to a container for storing and ejecting liquid (claims 1-13, 27-43), a method for manufacture of such container (claims 44-48, 52-53), a method for ejecting liquid from such container (claims 54-63, 92), a device for ejecting liquid from such container (claims 90-91).

.../...

1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:

4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

The additional search fees were accompanied by the applicant's protest.
 No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

International application No.
PCT/SE00/01514

Box II

Invention 2

Claims 14-26, 49-51. Relates to connected containers forming a multipel container unit (claims 14-26), a method for manufacture of such multipel container unit (claims 49-51).

Invention 3

Claims 64-89. Relates to a device for ejecting liquid from a container.

INTERNATIONAL SEARCH REPORT
Information on patent family members

03/10/00

International application No.
PCT/SE 00/01514

EP	0832568	A2	01/04/98	AT	163505	T	15/03/98
				AU	687372	B	26/02/98
				AU	5705894	A	04/07/94
				CA	2150719	A	23/06/94
				DE	69317301	D,T	17/09/98
				DK	674482	T	28/12/98
				EP	0674482	A,B	04/10/95
				SE	0674482	T3	
				EP	0919134	A	02/06/99
				ES	2115206	T	16/06/98
				FI	952980	A	16/06/95
				GB	9226307	D	00/00/00
				GR	3026653	T	31/07/98
				HK	1009636	A	00/00/00
				JP	8508157	T	03/09/96
				NO	952420	A	16/06/95
				NZ	258954	A	24/02/97
				US	5893485	A	13/04/99
				WO	9413154	A	23/06/94
				GB	9305018	D	00/00/00
EP	0139855	A1	08/05/85	DE	3324780	A	17/01/85
				JP	60075065	A	27/04/85
US	4433797	A	28/02/84	CA	1129384	A	10/08/82
				DE	2920497	A,C	27/11/80
				FR	2457231	A,B	19/12/80
				GB	2050303	A,B	07/01/81
				DE	2946814	A	27/05/81
US	5709202	A	20/01/98	AU	690561	B	30/04/98
				AU	708140	B	29/07/99
				AU	5829698	A	14/05/98
				AU	6956994	A	20/12/94
				CA	2162399	A	08/12/94
				EP	0701457	A	20/03/96
				JP	9503723	T	15/04/97
				US	5497763	A	12/03/96
				US	5544646	A	13/08/96
				US	5660166	A	26/08/97
				US	5718222	A	17/02/98
				US	5792057	A	11/08/98
				US	5823178	A	20/10/98
				US	6014969	A	18/01/00
				US	6123068	A	26/09/00
				WO	9427653	A	08/12/94
US	5516008	A	14/05/96	US	5425480	A	20/06/95

INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No.

03/10/00

PCT/SE 00/01514

WO	9606581	A1	07/03/96	AT	171858	T	15/10/98
				AU	703399	B	25/03/99
				AU	3391495	A	22/03/96
				BG	101365	A	30/12/97
				BR	9508672	A	28/10/97
				CA	2198892	A	07/03/96
				CZ	9700596	A	15/07/98
				DE	69505266	D, T	06/05/99
				DK	778758	T	21/06/99
				EP	0778758	A, B	18/06/97
				SE	0778758	T3	
				EP	0845253	A	03/06/98
				ES	2125654	T	01/03/99
				FI	970706	A	07/04/97
				GB	2306902	A	14/05/97
				GB	9417399	D	00/00/00
				GB	9704403	D	00/00/00
				HU	77167	A	02/03/98
				JP	10506028	T	16/06/98
				NO	970874	A	29/04/97
				NZ	292195	A	24/09/98
				PL	318935	A	21/07/97
				GB	9505472	D	00/00/00
				WO	9600050	A	04/01/96
				GB	9505474	D	00/00/00

WO	9723177	A1	03/07/97	AU	1201297	A	17/07/97
				CA	2240322	A	03/07/97
				EP	0873096	A	28/10/98
				GB	9526150	D	00/00/00
				JP	2000505663	T	16/05/00

GB	2255918	A	25/11/92	GB	9109064	D	00/00/00
				GB	9208178	D	00/00/00

US	4522622	A	11/06/85	NONE			
----	---------	---	----------	------	--	--	--

INTERNATIONAL SEARCH REPORT

Information on patent family members

03/10/00

International application No.
PCT/SE 00/01514

WO	9600050	A1	04/01/96	GB	9412665	D	00/00/00
				AT	171858	T	15/10/98
				AU	703399	B	25/03/99
				AU	3391495	A	22/03/96
				BG	101365	A	30/12/97
				BR	9508672	A	28/10/97
				CA	2198892	A	07/03/96
				CZ	9700596	A	15/07/98
				DE	69505266	D,T	06/05/99
				DK	778758	T	21/06/99
				EP	0778758	A,B	18/06/97
				SE	0778758	T3	
				EP	0845253	A	03/06/98
				ES	2125654	T	01/03/99
				FI	970706	A	07/04/97
				GB	2306902	A	14/05/97
				GB	9505472	D	00/00/00
				GB	9704403	D	00/00/00
				HU	77167	A	02/03/98
				JP	10506028	T	16/06/98
				NO	970874	A	29/04/97
				NZ	292195	A	24/09/98
				PL	318935	A	21/07/97
				WO	9606581	A	07/03/96

US	1751341	A	18/03/30	NONE
----	---------	---	----------	------

US	2649995	A	25/08/53	NONE
----	---------	---	----------	------

US	4236516	A	02/12/80	US	4325369	A	20/04/82
----	---------	---	----------	----	---------	---	----------

US	5429280	A	04/07/95	DE	69301974	D,T	28/11/96
				EP	0581947	A,B	09/02/94
				SE	0581947	T3	
				ES	2085769	T	01/06/96
				FR	2687643	A,B	27/08/93
				JP	6507140	T	11/08/94
				US	5520310	A	28/05/96
				WO	9316941	A	02/09/93
				DE	69321703	D,T	18/03/99
				EP	0610640	A,B	17/08/94
				FR	2700404	A,B	13/07/94
				US	5397885	A	14/03/95

INTERNATIONAL SEARCH REPORT

Information on patent family members

03/10/00

International application No.

PCT/SE 00/01514

US	4072249	A	07/02/78	CA	1047459	A	30/01/79
				CH	605328	A	29/09/78
				DE	2608794	A	16/09/76
				DK	84076	A	04/09/76
				DK	141895	B,C	14/07/80
				FR	2302936	A,B	01/10/76
				GB	1539598	A	31/01/79
				IN	145108	A	26/08/78
				JP	51143481	A	09/12/76
				NL	7602203	A	07/09/76
				SE	7502318	A	06/09/76
				US	4282986	A	11/08/81

GB	2246555	A	05/02/92	GB	9015905	D	00/00/00
----	---------	---	----------	----	---------	---	----------

US	5622166	A	22/04/97	AU	695051	B	06/08/98
				AU	5251096	A	18/11/96
				BG	101992	A	31/07/98
				BR	9608194	A	21/07/98
				CA	2217672	A	31/10/96
				CN	1181709	A	13/05/98
				CZ	9703349	A	14/01/98
				EP	0835147	A	15/04/98
				HU	9801656	A	28/10/98
				IL	117706	D	00/00/00
				JP	10512788	T	08/12/98
				NO	974910	A	24/10/97
				NZ	304654	A	28/01/99
				PL	322909	A	02/03/98
				RO	115407	A	28/02/00
				SK	144697	A	04/03/98
				US	5921237	A	13/07/99
				US	6029663	A	29/02/00
				WO	9633759	A	31/10/96

US	5349947	A	27/09/94	EP	0614460	A	14/09/94
				JP	6510560	T	24/11/94

US	5415162	A	16/05/95	NONE			
----	---------	---	----------	------	--	--	--

INTERNATIONAL SEARCH REPORT

Information on patent family members

International application No.

03/10/00

PCT/SE 00/01514

US	5221050	A	22/06/93	AT	179097	T	15/05/99
				CA	2054224	A	27/04/92
				DE	69131153	D,T	18/11/99
				DK	482814	T	01/11/99
				EP	0482814	A,B	29/04/92
				SE	0482814	T3	
				ES	2133280	T	16/09/99
				GB	9023339	D	00/00/00
				GB	9206998	D	00/00/00
				GR	3030738	T	30/11/99
				JP	5138081	A	01/06/93